

ence Range: 1 to 6200

```

      10      20      30      40      50      60      70
GACGGATCGG GAGATCTCCC GATCCCCTAT GGTGACTCTT CAGTACAATC TGCTCTGATG CCGCATAGTT

      80      90     100     110     120     130     140
AAGCCAGTAT CTGCTCCCTG CTTGTGTGTT GGAGGTCGCT GAGTAGTGGC CGAGCAAAAT TTAAGCTACA

      150     160     170     180     190     200     210
ACAAAGCAAG GCTTGACCGA CAATTGCATG AAGAATCTGC TTAGGGTTAG GCGTTTTGCG CTGCTTCGCG

      220     230     240     250     260     270     280
ATGTACGGGC CAGATATACG CGTTGACATT GATTATTGAC TAGTTATTAA TAGTAATCAA TTACGGGGTC

      290     300     310     320     330     340     350
ATTAGTTCAT AGCCCATATA TGGAGTTCCG CGTTACATAA CTTACGTAA ATGGCCCGCC TGGCTGACCG

      360     370     380     390     400     410     420
CCCAACGACC CCCGCCATT GACGTCAATA ATGACGTATG TTCCCATAGT AACGCCAATA GGGACTTTCC

      430     440     450     460     470     480     490
ATTGACGTCA ATGGGTGGAC TATTTACGGT AAATGCCCCA CTGGGCAGTA CATCAAGTGT ATCATATGCC

      500     510     520     530     540     550     560
AAGTACGCCC CCTATTGACG TCAATGACGG TAAATGGCCC GCCTGGCATT ATGCCCAGTA CATGACCTTA

      570     580     590     600     610     620     630
TGGGACTTTC CTACTTGCCA GTACATCTAC GTATTAGTCA TCGCTATTAC CATGGTGATG CGGTTTGGC

      640     650     660     670     680     690     700
AGTACATCAA TGGGCGTGA TAGCGGTTTG ACTCAGGGGG ATTTCCAAGT CTCCACCCCA TTGACGTCAA

      710     720     730     740     750     760     770
TGGGAGTTTG TTTTGGCACC AAAATCAACG GGACTTTCCA AAATGTCGTA ACAACTCCGC CCCATTGACG

      780     790     800     810     820     830     840
CAATGGGGCG GTAGCGGTGT ACGGTGGGAG GTCTATATAA GCAGAGCTCT CTGGCTAACT AGAGAACCCA

      850     860     870     880     890     900     910
CTGCTTAAGT GGCTTATCGA AATTAATACG ACTCACTATA GGTAGACCCA AGCTTCGCAG AATTCCTGCG

      920     930     940     950     960     970     980
GCTGCTACAG TGTGTCCAGC GTCCTGCCTG GCTGTGCTGA GUGCTGGAAC AGTGGCGCAT CATTCAGTGT

      990    1000    1010    1020    1030    1040    1050
CACAGTTACC CATCTGAGT CTGGCACCTT AACTGGCACA ATTGCCAAGG TCACAGGTGA GCTCAGATGC

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0993241.111001

FIGURE 1

090341-1101

P. 31 (sup'd)

FIGURE 1A

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      _ _ _ _ _ ORF RF[1] _ _ _ _ _
    *   *   *   *   *   *   *   *   *   *   *   *   *   *   *
      1890       1900       1910       1920       1930       1940
AAC TTC ACC CAC CTG GAC GGG TAC CTG GAC CTT CTC AGG GAG AAC CAG CTC CTC CCA
Asn Phe Thr His Leu Asp Gly Tyr Leu Asp Leu Leu Arg Glu Asn Gln Leu Leu Pro>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

      1950       1960       1970       1980       1990
GGG TTT GAG CTG ATG GGC AGC GCC TCG GGC CAC TTC ACT GAC TTT GAG GAC AAG CAG
Gly Phe Glu Leu Met Gly Ser Ala Ser Gly His Phe Thr Asp Phe Glu Asp Lys Gln>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

2000       2010       2020       2030       2040       2050
CAG GTG TTT GAG TGG AAG GAC TTG GTC TCC AGC CTG GCC AGG AGA TAC ATC GGT AGG
Gln Val Phe Glu Trp Lys Asp Leu Val Ser Ser Leu Ala Arg Arg Tyr Ile Gly Arg>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

      2060       2070       2080       2090       2100       2110
TAC GGA CTG GCG CAT GTT TCC AAG TGG AAC TTC GAG ACG TGG AAT GAG CCA-GAC CAC
Tyr Gly Leu Ala His Val Ser Lys Trp Asn Phe Glu Thr Trp Asn Glu Pro Asp His>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

      2120       2130       2140       2150       2160
CAC GAC TTT GAC AAC GTC TCC ATG ACC ATG CAA GGC TTC CTG AAC TAC TAC GAT GCC
His Asp Phe Asp Asn Val Ser Met Thr Met Gln Gly Phe Leu Asn Tyr Tyr Asp Ala>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

2170       2180       2190       2200       2210       2220
TGC TCG GAG GGT CTG CGC GCC GCC AGC CCC GCC CTG CGG CTG GGA GGC CCC GGC GAC
Cys Ser Glu Gly Leu Arg Ala Ala Ser Pro Ala Leu Arg Leu Gly Gly Pro Gly Asp>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

      2230       2240       2250       2260       2270       2280
TCC TTC CAC ACC CCA CCG CGA TCC CCG CTG AGC TGG GGC CTC CTG CGC CAC TGC CAC
Ser Phe His Thr Pro Pro Arg Ser Pro Leu Ser Trp Gly Leu Leu Arg His Cys His>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

      2290       2300       2310       2320       2330       2340
GAC GGT ACC AAC TTC TTC ACT GGG GAG GCG GGC GTG CGG CTG GAC TAC ATC TCC CTC
Asp Gly Thr Asn Phe Phe Thr Gly Glu Ala Gly Val Arg Leu Asp Tyr Ile Ser Leu>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

      2350       2360       2370       2380       2390
CAC AGG AAG GST GCG CGC AGC TCC ATC TCC ATC CTG GAG CAG GAG AAG GTC GTC GCG
His Arg Lys Gly Ala Arg Ser Ser Ile Ser Ile Leu Glu Gln Glu Lys Val Val Ala>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

2400       2410       2420       2430       2440       2450
CAG CAG ATC CGG CAG CTC TTC CCC AAG TTC GCG GAC ACC CCC ATT TAC AAC GAC GAG
Gln Gln Ile Arg Gln Leu Phe Pro Lys Phe Ala Asp Thr Pro Ile Tyr Asn Asp Glu>
_ _ _ _ _ ORF RF[1] _ _ _ _ _

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FIGURE 1B

0993241.11701

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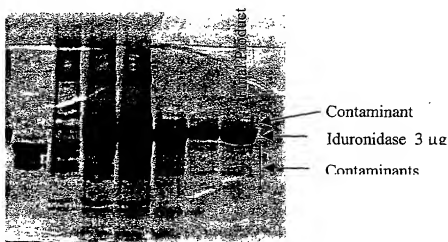
AAAAAAAAA AAAAAAAAAAG AATTCTCGCA GCCCGGGGGA TCCACTAGTT CTAGAGGGCC CGTTTAAACC
  3760      3770      3780      3790      3800      3810      3820
*          *          *          *          *          *          *
CGCTGATCAG CCTCGACTGT GCCTTCTAGT TGCCAGCCAT CTGTTGTTTG CCCCTCCCCC GTGCCTTCCT
  3830      3840      3850      3860      3870      3880      3890
*          *          *          *          *          *          *
TGACCCTGGA AGGTGCCACT CCCACTGTCC TTTCCTAATA AAATGAGGAA ATTGCATCGC ATTGCTGAG
  3900      3910      3920      3930      3940      3950      3960
*          *          *          *          *          *          *
TAGGTGTCAT TCTATTCTGG GGGGTGGGGT GGGGCAGGAC AGCAAGGGGG AGGATTGGGA AGACAATAGC
  3970      3980      3990      4000      4010      4020      4030
*          *          *          *          *          *          *
AGGCATGCTG GGGATGCGGT GGGCTCTATG GCTTCTGAGG CGGAAGAAGC CAGCTGGGGC TCGAGAGCTT
  4040      4050      4060      4070      4080      4090      4100
*          *          *          *          *          *          *
GGCGTAATCA TGGTCATAGC TGTTTCCTGT GTGAAATTGT TATCCGCTCA CAATTCCACA CAACATACGA
  4110      4120      4130      4140      4150      4160      4170
*          *          *          *          *          *          *
GCCGGAAGCA TAAAGTGTA AGCCTGGGGT GCCTAATGAG TGAGCTAACT CACATTAATT GCGTTGCGCT
  4180      4190      4200      4210      4220      4230      4240
*          *          *          *          *          *          *
CACTGCCCGC TTTCCAGTCG GGAACCTGT CGTGCCAGCT GCATTAATGA ATCGGCCAAC GCGCGGGGAG
  4250      4260      4270      4280      4290      4300      4310
*          *          *          *          *          *          *
AGCGGGTTTG CGTATTGGGC GCTCTCCGC TTCTCGCTC ACTGACTCGC TCGCTCGGT CGTTCGGGTG
  4320      4330      4340      4350      4360      4370      4380
*          *          *          *          *          *          *
CGCGAGCGG TATCAGTCA CTCAAAGGCG GTRATACGTT TATCCACAGA ATCAGGGGAT AACGCAGGAA
  4390      4400      4410      4420      4430      4440      4450
*          *          *          *          *          *          *
AGAACATGTG AGCAAAAGGC CAGCAAAAGG CCAGGAACCG TAAAAAGGCC GCGTTGCTGG CGTTTTTCCA
  4460      4470      4480      4490      4500      4510      4520
*          *          *          *          *          *          *
TAGGCTCCGC CCCCTGAGC AGCATCACAA AAATCGAGCG TCAAGTCAGA GGTGGCGAAA CCCGACAGGA
  4530      4540      4550      4560      4570      4580      4590
*          *          *          *          *          *          *
CTATAAAGAT ACCAGGCGTT TCCCCTGGA AGCTCCCTCG TCGCTCTCC TGTTCGACC CTGCGCGTTA
  4600      4610      4620      4630      4640      4650      4660
*          *          *          *          *          *          *
CCGGATACCT GTCCGCTTT CTCCCTTCGG GAAGCGTGGC GCTTCTCAA TGCTCAGCT GTAGGTATCT
  4670      4680      4690      4700      4710      4720      4730
*          *          *          *          *          *          *
CAGTTCGGTG TAGGTCGTT GCTCCAAGCT GGGCTGTGTG CACGAACCCC CCGTTCAGCC CGACCGTGC
  4740      4750      4760      4770      4780      4790      4800
*          *          *          *          *          *          *
GCCTTATCCG GTAACATCG TCTTGAGTCC AACCCGGTAA GACACGACTT ATCGCACTG GCACAGCCCA

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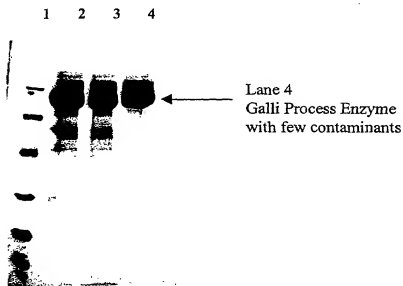
FIGURE 1E

FIGURE 2. SDS-POLYACRYLAMIDE GELS DEMONSTRATING IMPROVEMENTS IN PURITY

Gel using the Kakkis et al 1994, published procedure for purification



Gel using the new Galli Process contained in this application



1. Molecular Weight Marker
2. Prior Process Carson (nonpublished) Batch 2000C9001 Reference Reduced (7.5 µg)
3. Same Batch 2000C9001 Reference Reduced (5.0 µg)
4. Galli Process Enzyme Batch P10006 (5.0 µg)

FIGURE 2

FIGURE 3A IDURONIDASE PRODUCTION USING THE GALLI PROCESS

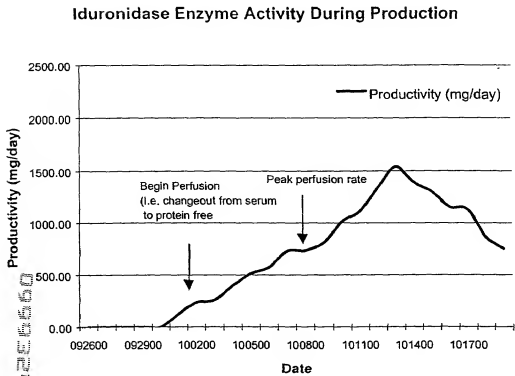
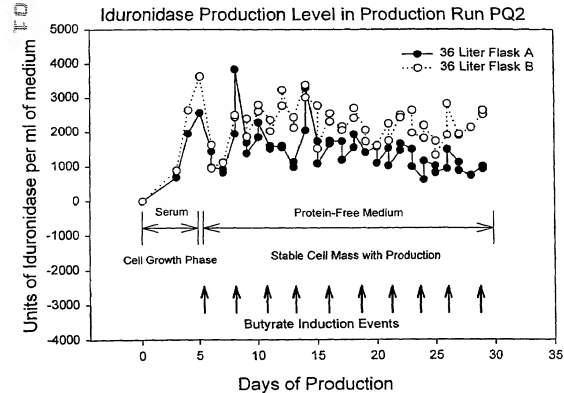


FIGURE 3B. IDURONIDASE PRODUCTION USING BUTYRATE INDUCTION



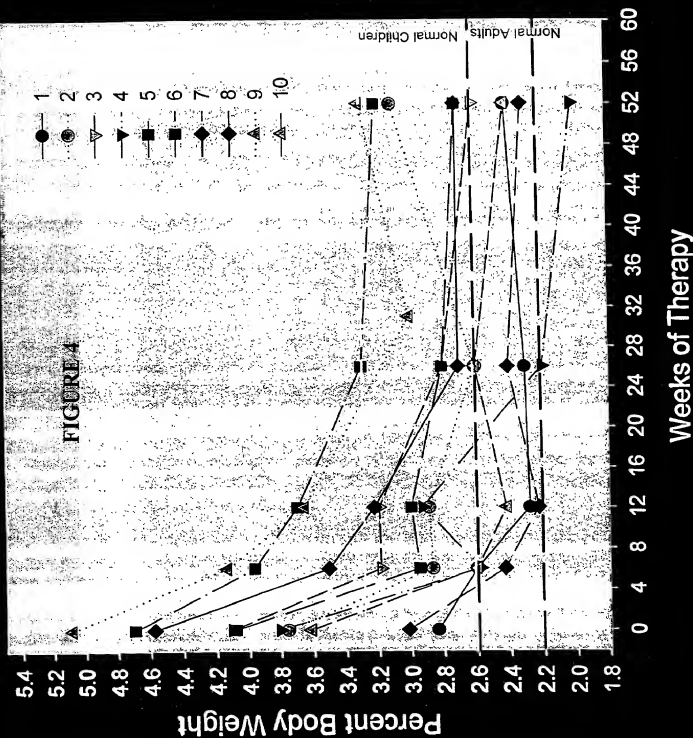


FIGURE 4

Urinary GAG Excretion During Enzyme Therapy

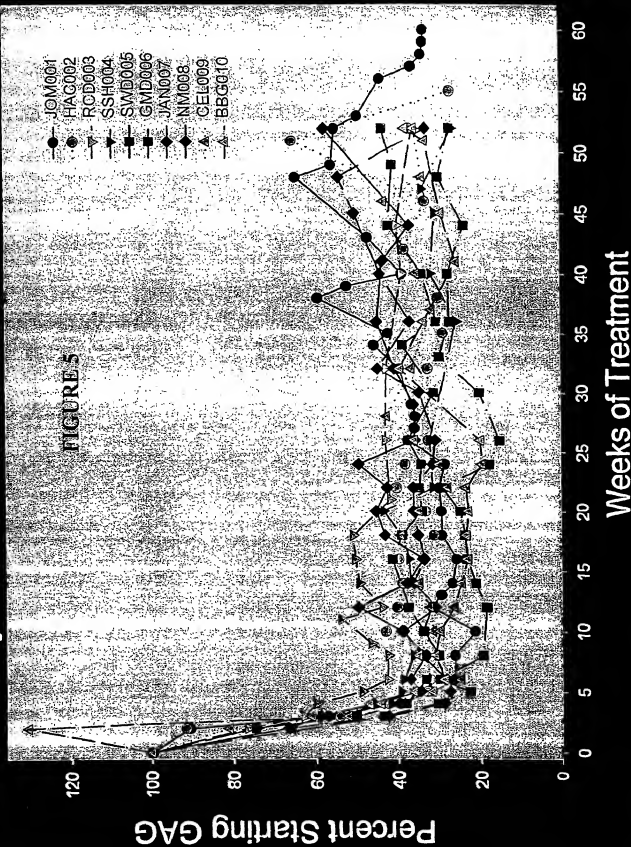


FIGURE 5

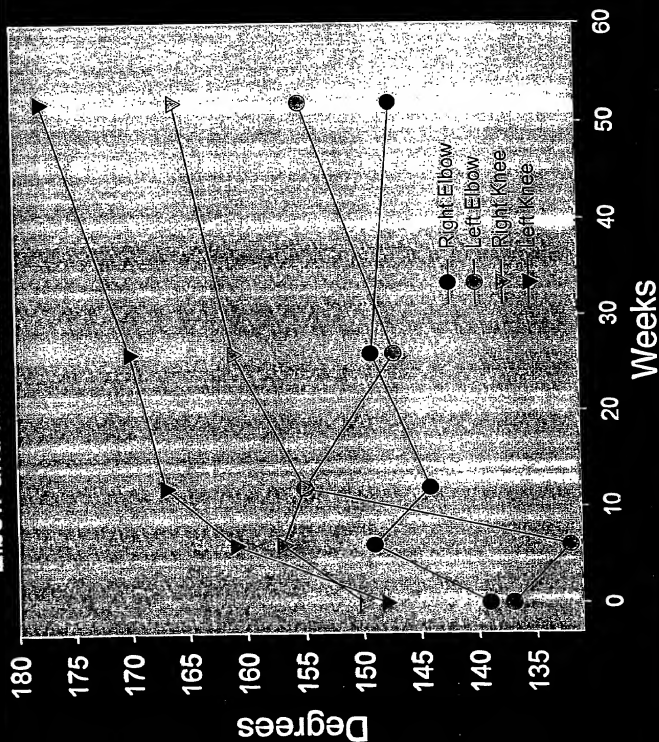


FIGURE 6

Snouder Extension
four patients with most restriction

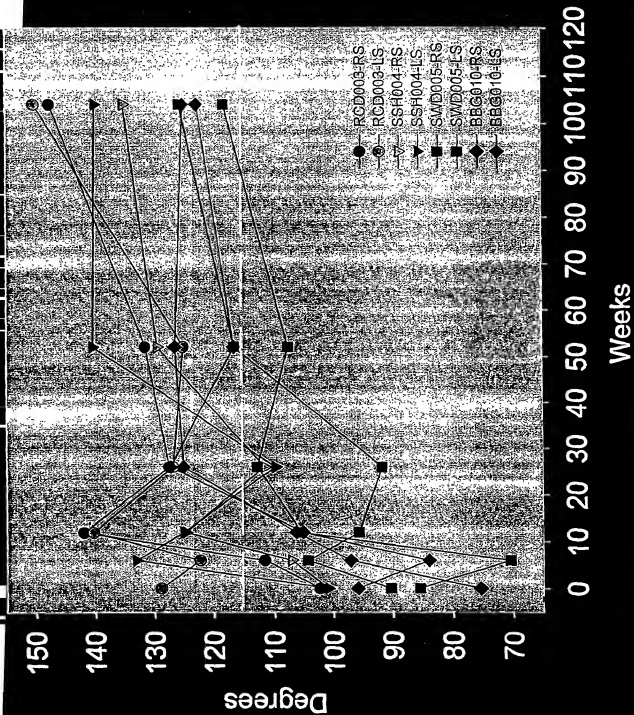


FIGURE 7

Sleep Apnea Improves

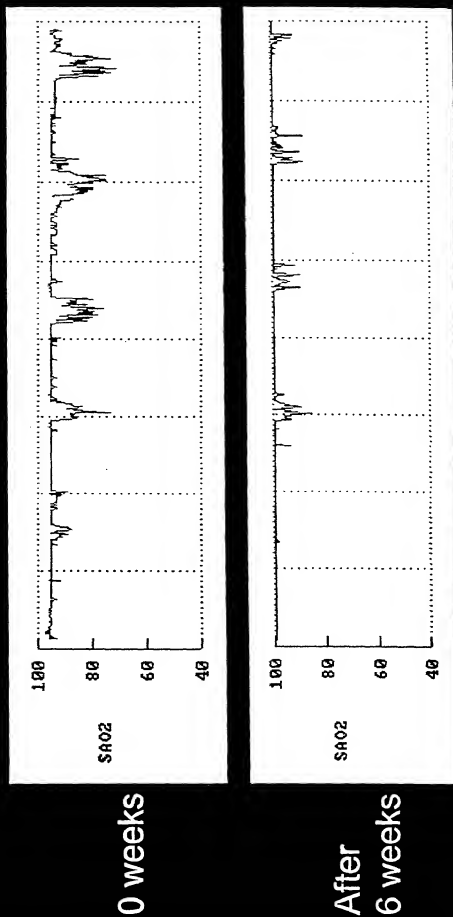


FIGURE 8

Apneas + Hypopneas During Sleep Pre and Post Treatment

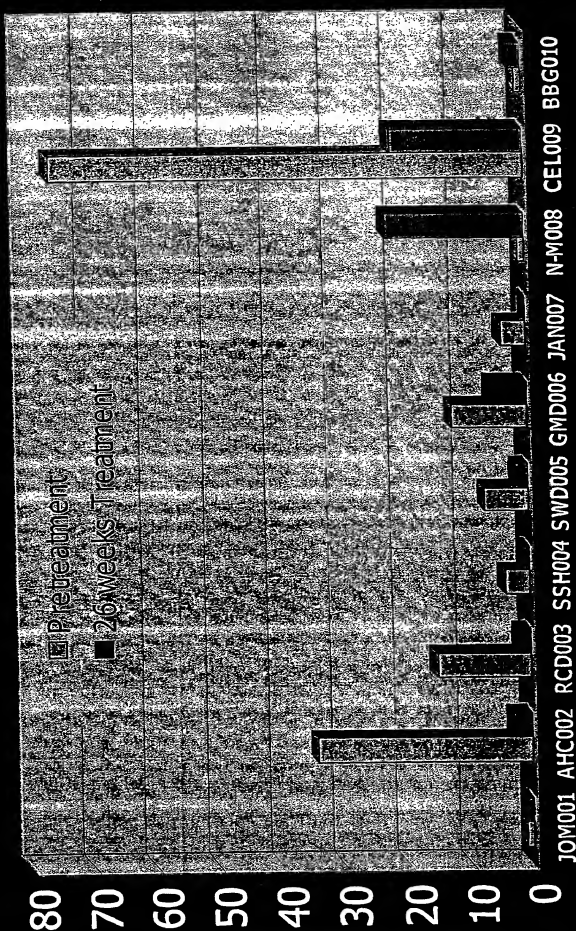


FIGURE 9

Pulmonary Function Tests in GIVD006

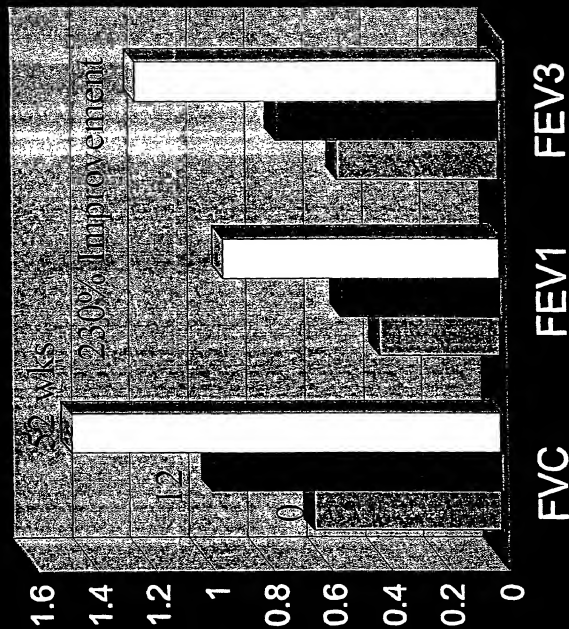


FIGURE 10

Increased Height Growth Velocity

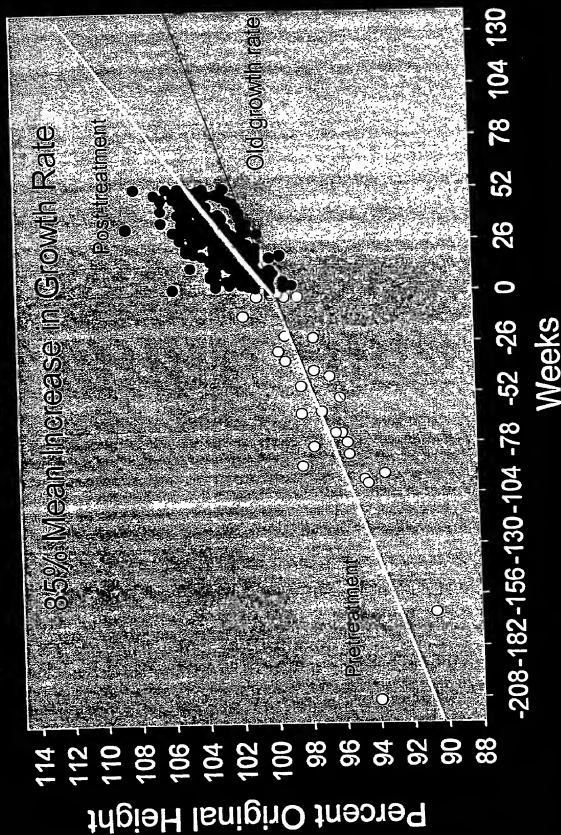


FIGURE 11

FIGURE 12.

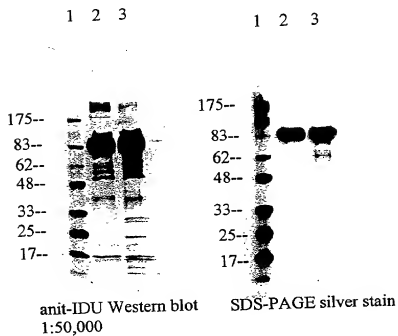
COMPARISON OF HOST PROTEIN CONTAMINATION BETWEEN A PRIOR AND THE NEW GALLI PROCESS

Chinese Hamster Ovary Host Protein Contamination by ELISA Assay

SOURCE AND BATCH NUMBER	CHOP PROTEIN CONTAMINATION (microgram per milligram)	PERCENT CHOP CONTAMINATION	PURITY OF THE ENZYME FROM CHOP
Prior Process (Carson/REI)			
C9002	14	1.4%	98.6%
C9003	24	2.4%	97.6%
C9004	16	1.6%	98.4%
New Process (Galli)			
P1003	<1.3	<0.13%	>99.9%
P1006	1.2	0.12%	99.9%
P1007	<0.6	<0.06%	>99.9%
P1008	<0.67	<0.067%	>99.9%

FIGURE 12

Comparison of Galli and Carson Material



- 1 Marker
- 2 Galli Referenced-0201
- 3 Carson C9002

5ug/lane

FIGURE 13